Title: Anti-TNF Antibodies and Peptides.. Inventors: Junming Le, *et al*.

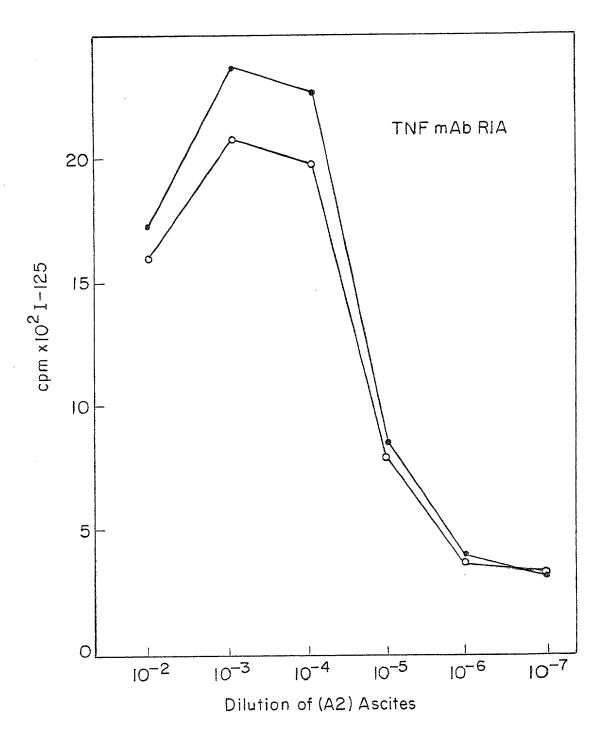


FIG. 1

Title: Anti-TNF Antibodies and Peptides...

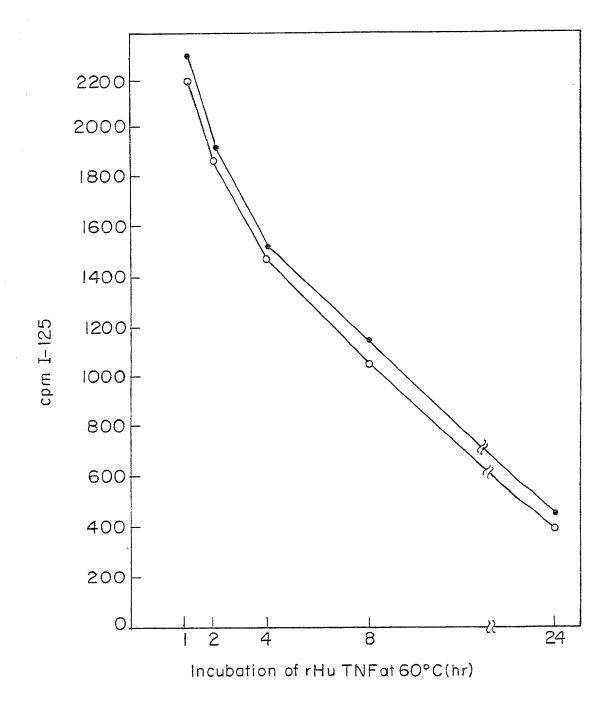
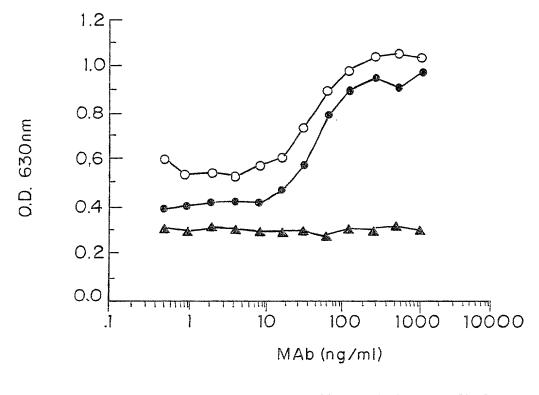


FIG. 2

0975.1005-010

Title: Anti-TNF Antibodies and Peptides...

Junming Le, et al. Inventors:

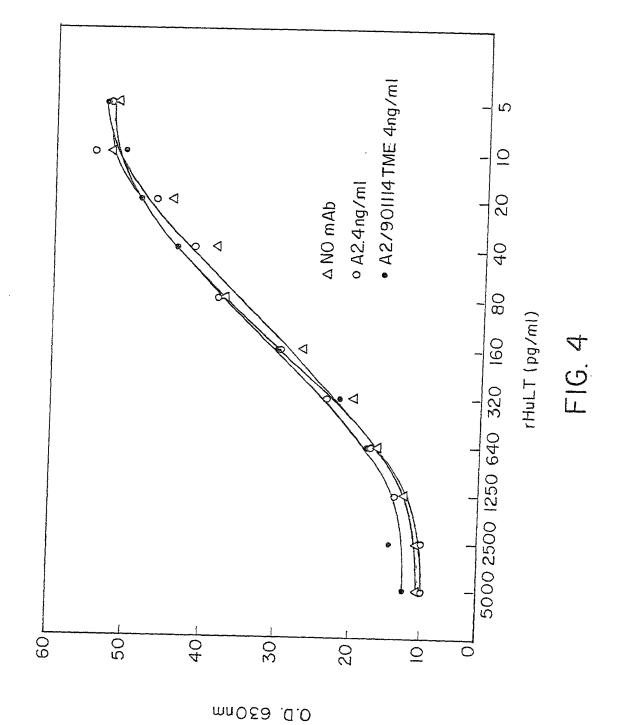


Natural Human TNF

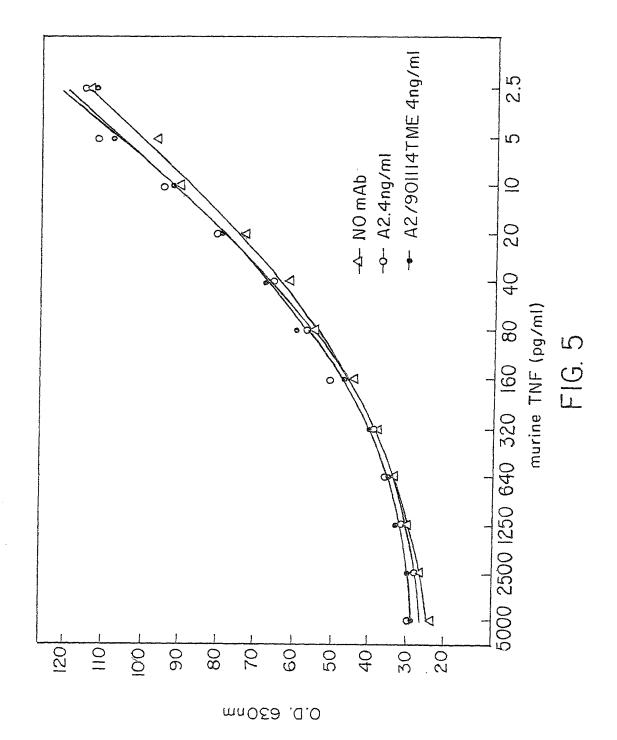
- Recombinant Human TNF 0
- Control

FIG. 3

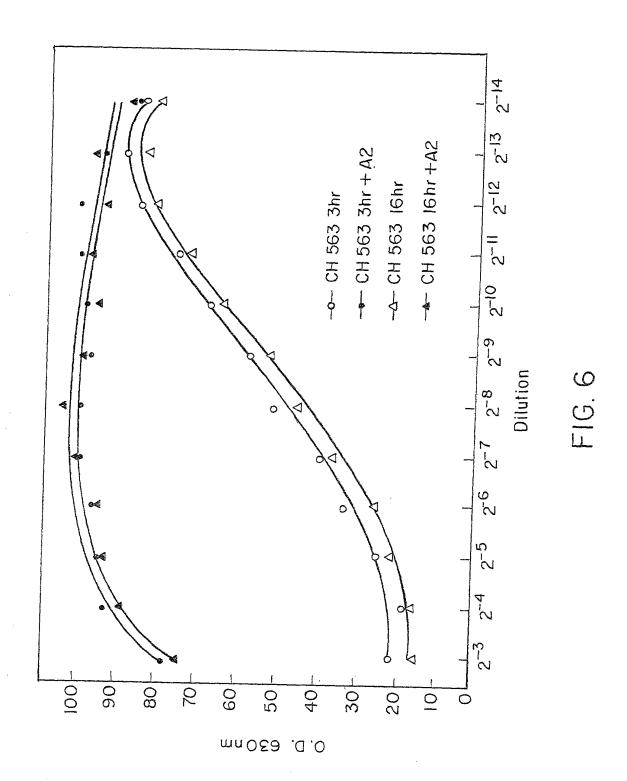
Title: Anti-TNF Antibodies and Peptides...



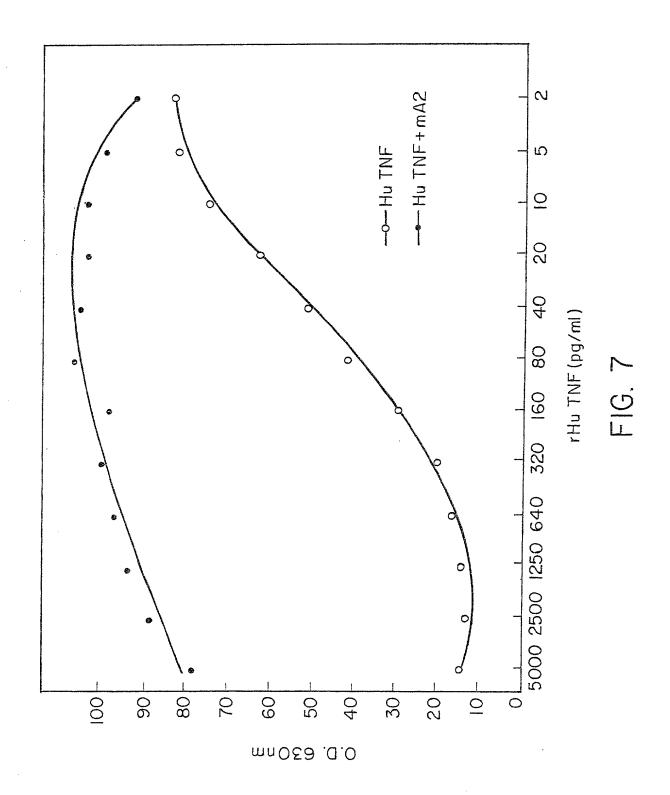
Title: Anti-TNF Antibodies and Peptides...



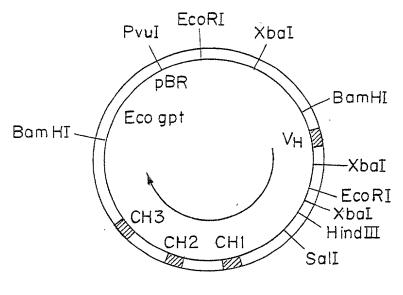
Title: Anti-TNF Antibodies and Peptides...



Docket No.: 0975.1005-010
Title: Anti-TNF Antibodies and Peptides...
Inventors: Junming Le, et al.



Title: Anti-TNF Antibodies and Peptides...



pA2HG1apgpt

FIG. 8A

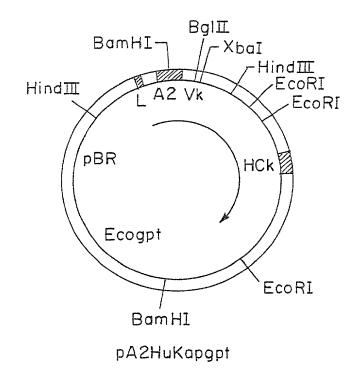


FIG. 8B

Title: Anti-TNF Antibodies and Peptides...

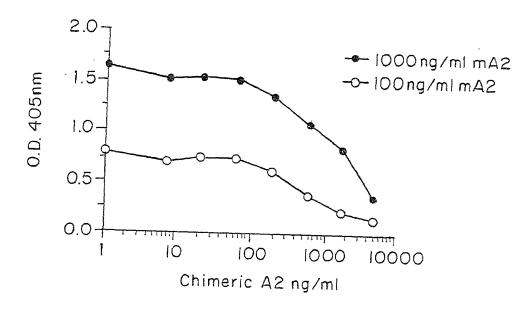


FIG. 9A

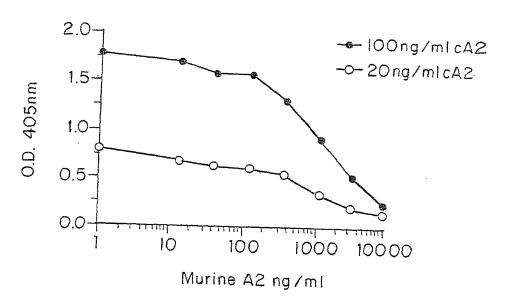


FIG. 9B

0975.1005-010 Docket No.:

Title: Anti-TNF Antibodies and Peptides...

Junming Le, et al. Inventors:

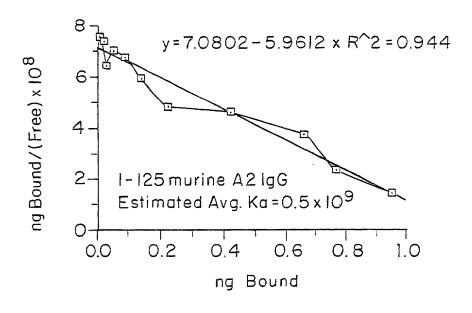


FIG. 10A

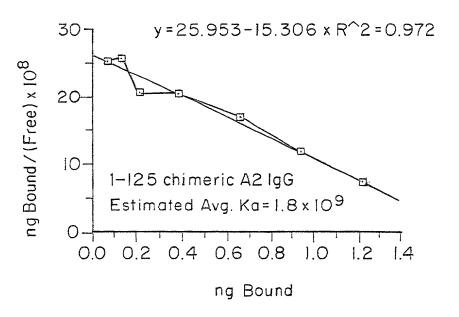


FIG. 10B

Title: Anti-TNF Antibodies and Peptides...

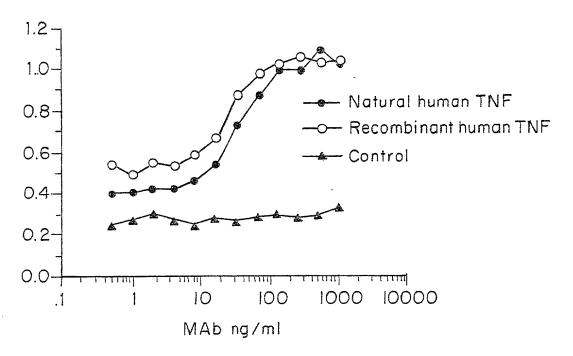


FIG. 11

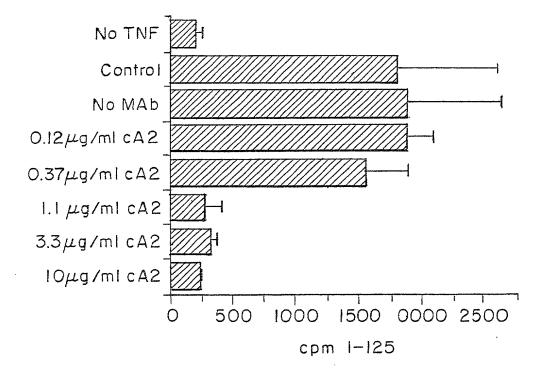


FIG. 12

Title: Anti-TNF Antibodies and Peptides... Inventors: Junming Le, *et al*.

| Pro | ${\tt Gly}$ | Ser | Ile | Pro | Leu | Asp | |
|----------|-------------|------------------|----------------------|-----------------------------------|-------------------------|--------------------|-----------------|
| Asn | Asn | Tyr | Thr | Ser | Tyr | Pro | |
| Ala | Ala | | | Lys | Ile | Arg | |
| Val | Leu | Leu Ile | Thr His | Ile | Pro | Asn | Leu |
| Val | Leu | | Leu | | Glu | Ile Asn | Ala |
| His | Ala Leu | Glu Gly Leu Tyr | Leu | 90 Lys Val Asn Leu Leu Ser Ala | $\mathrm{T}\mathrm{yr}$ | Glu | Ile |
| Ala | | ${	t G1} {	t y}$ | | Leu | Pro Trp Tyr | Leu Ser Ala | Phe Gly Ile |
| Val | Ala Asn | Glu | His Val | Leu | Pro | Ser | G1y |
| Pro | Arg | Ser | \mathtt{Thr} | Asn | Lys | Leu | Phe |
| Lys | Arg | Pro | Ser | Val | 110 Glu Ala | 130 Asp Arg | 150 Val Tyr |
| Asp | 30 Asn | 50 Val | 70 Pro | 90 Lys | 110 Glu | | 150 Val |
| Ser | Trp Leu | Leu Val | $Cy\mathtt{s}$ | Thr | Ala | \mathtt{Gly} | Gly Gln |
| Pro | Trp | | Gly Gln Gly Cys | Gln | Pro Glu Gly Ala | Lys | ${	t Gly}$ |
| Arg Thr | Gln | Asn Gln | Gln | Ser Tyr | Glu | Glu | Ser |
| Arg | Leu | | | | | Leu | Glu |
| Ser | Gln | Asp | Lys | Val | Thr | Gln | Ala |
| Ser | Glu Gly Gln | Leu Arg | Phe | Ala | Gln Arg Glu | Phe | Leu Asp Phe Ala |
| Ser | Glu | Leu | Leu | H H | . Arg | . Val | Asp |
| Arg | Ala | Glu | Val | . Arg | | 121 Gly Gly Val | Leu |
| l Val | 21 Gln | 41 Val | 61 Gln | 81 Ser | 101 Cys | 121 Gly | 141 Tyr |
| | | | | | | | |

Inventors:

Junming Le, et al.

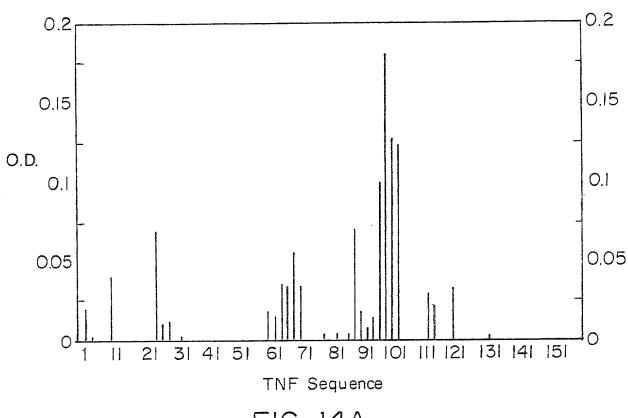


FIG. 14A

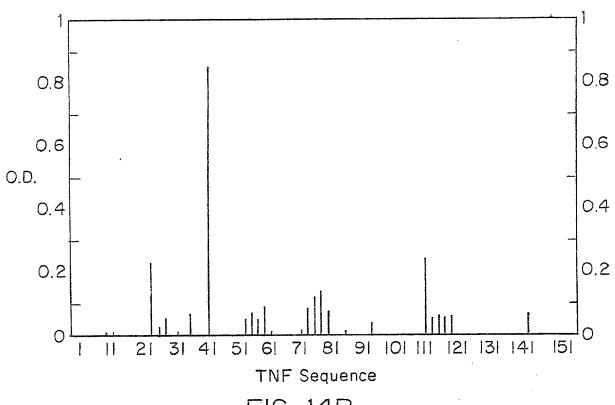


FIG. 14B

Docket No.: 0975.1005-010

Title: Anti-TNF Antibodies and Peptides...

Inventors:

Junming Le, et al.

Pro

Ser

Lys

Ala

Ser

90 Tyr Gln Thr Lys Val Asn Leu Leu

Ser

Ala Val

Arg

81 Ser

| | Ç. |
|---|---|
| Pro | G1y |
| 10 Thr Pro Ser Asp Lys Pro Val Ala His Val Val Ala Asn Pro | n Arg Arg Ala Asn Ala Leu Leu Ala Asn Gly |
| Ala | Ala |
| Val | Leu |
| Val | Leu |
| His | Ala |
| Ala | Asn |
| Val | Ala |
| Pro | Arg |
| Lys | Arg |
| 10 Asp | 30 Gln Trp Leu Asn |
| Ser | Leu |
| Pro | Trp |
| \mathtt{Thr} | Gln |
| Arg | Leu |
| Ser Ser | Ala Glu Gly Gln |
| Ser | Gly |
| Ser | Glu |
| 1 Val Arg Ser | |
| 1 Val | 21 G1n |

| Ser | Ile |
|-----------------------------------|-----------------------|
| Tyr | Thr |
| Glu Gly Leu Tyr Leu Ile Tyr Ser | His |
| Leu | Thr |
| ľyr | Leu |
| ren ' | Leu Leu Thr |
| GΙΥ | Val |
| g]u | Thr His |
| Ser | Thr |
| Pro Ser | Ser |
| 50 Val | 70 Pro |
| Leu Val | 70 Jln Gly Cys Pro |
| Leu | Gly |
| Gln | Gln |
| Asn | Lys Gly Glı |
| Asp | Lys |
| Arg | Phe |
| 41 Val Glu Leu Arg Asp Asn Gln | . Val Leu Phe |
| Glu | Val |
| 41 Val | 61 G1n |

| | ī |
|--|--------------------------------------|
| | Ile Tyr Leu |
| | Tyr |
| | Ile |
| | ro |
| | 31u |
| | lyr (|
| | 110 Glu Ala Lys Pro Trp Tyr Glu P |
| | ro 1 |
| | ys F |
| | la L |
| | 110 Glu A |
| | la G |
| | Gly Ala (|
| | Ħ |
| İ | o G] |
| | Thr Pro Gl |
| | Th |
| | G1 v |
| | Arg |
| market supported to the control of t | Gln |
| | 101 Cys Gln Arg Glu T |
| | |

| | _ |
|-----|---------|
| | Asp |
| | Pro |
| | Arg |
| | Asn . |
| | Ile |
| | Glu : |
| | Ala G |
| | |
| | Se |
| | Leu Ser |
| | Arg |
| 130 | Asp |
| | Gl Y |
| | Lys |
| | Glu |
| | Leu |
| | Gln |
| | Phe |
| | Val |
| | Gly |
| 121 | Gly G |
| 1 1 |) |

Tyr Phe Gly Ile 141 Tyr Leu Asp Phe Ala Glu Ser Gly Gln Val

Title: Anti-TNF Antibodies and Peptides...

Inventors: Junming Le, et al.

GACATCTTGCTGACTCAGTCTCCAGCCATCCTGTCTGTGAGTCCAGGAGAAAGAGTCAGT ${\tt AspIleLeuLeuThrGlnSerProAlaIleLeuSerValSerProGlyGluArgValSer}$

PheserCysArgAlaSerGlnPheValGlySerSerIleHisTrpTyrGlnGlnArgThr TTCTCCTGCAGGCCCAGTCAGTTCGTTGGCTCAAGCATCCACTGGTATCAGCAAAGAACA

AATGGTTCTCCAAGGCTTCTCATAAAGTATGCTTCTGAGTCTATGTCTGGGATCCCTTCC ${\tt AsnGlySerProArgLeuLeuIleLysTyrAlaSerGluSerMetSerGlyIleProSer}$

 ${\tt ArgPheSerGlySerGlySerGlyThrAspPheThrLeuSerIleAsnThrValGluSer}$ AGGTTTAGTGGCAGTGGATCAGGGACAGATTTTACTCTTAGCATCAACACTGTGGAGTCT

GAAGATATTGCAGATTATTACTGTCAAGAAAGTCATAGCTGGCCATTCACGTTCGGCTCG ${\tt GluAspIleAlaAspTyrTyrCysGlnGlnSerHisSerTrpProPheThrPheGlySer}$

GGGACAAATTTGGAAGTAAAA GlyThrAsnLeuGluValLys FIG. 16A

Title: Anti-TNF Antibodies and Peptides...

Inventors: Junming Le, et al.

GAAGTGAAGCTTGAGGAGTCTGGAGGGCTTGGTGCAACCTGGAGGATCCATGAAACTC GluValLysLeuGluGluSerGlyGlyGlyLeuValGlnProGlyGlySerMetLysLeu TCCTGTGTTTCCTTCTTTTTTCAGTAACCACTGGATGAACTGGGTCCGCCAGTCT SerCysValAlaSerGlyPheIlePheSerAsnHisTrpMetAsnTrpValArgGlnSer CCAGAGAAGGGGCTTGAGTTGCTGAAATTAGATCAAAATCTATTAATTCTGCAACA $\mathtt{ProGluLysGlyLeuGluTrpValAlaGluIleArgSerLysSerIleAsnSerAlaThr}$

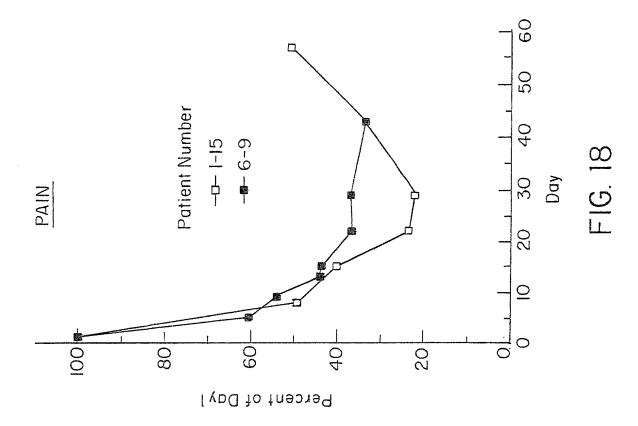
CATTATGCGGAGTCTGTGAAAGGGAGGTTCACCATCTCAAGAGATGATTCCAAAAGTGCT HisTyrAlaGluSerValLysGlyArgPheThrIleSerArgAspAspSerLysSerAla

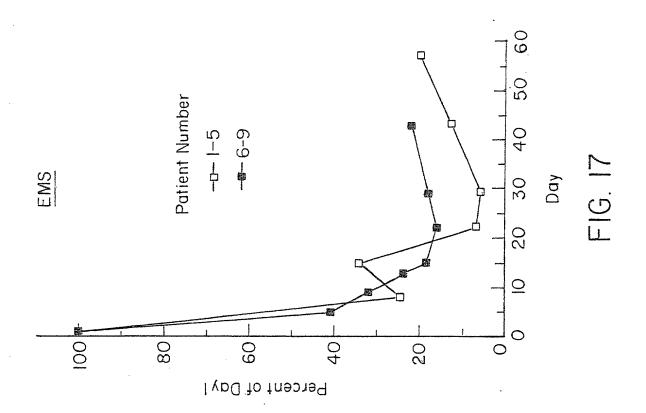
GTGTACCTGCAAATGACCGACTTAAGAACTGAAGACACTGGCGTTTTATTACTGTTCCAGG ${\tt ValTyrLeuGlnMetThrAspLeuArgThrGluAspThrGlyValTyrTyrCysSerArg}$

AATTACTACGGTAGTACCTACGACTACTGGGGCCAAGGCACCACTCTCACAGTGTCC AsnTyrTyrGlySerThrTyrAspTyrTrpGlyGlnGlyThrThrLeuThrValSer

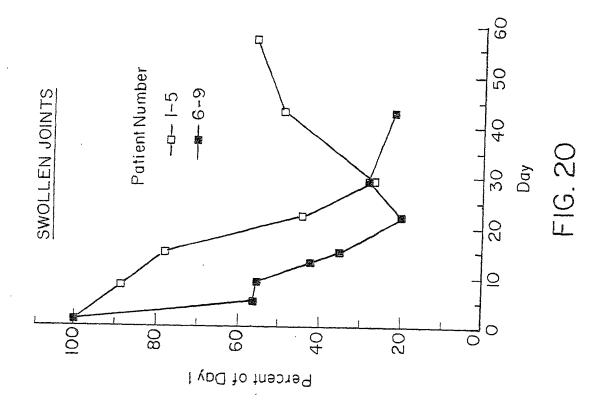
FIG. 16E

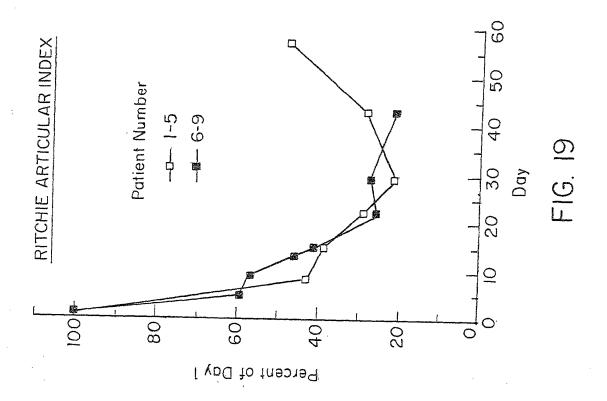
Title: Anti-TNF Antibodies and Peptides...



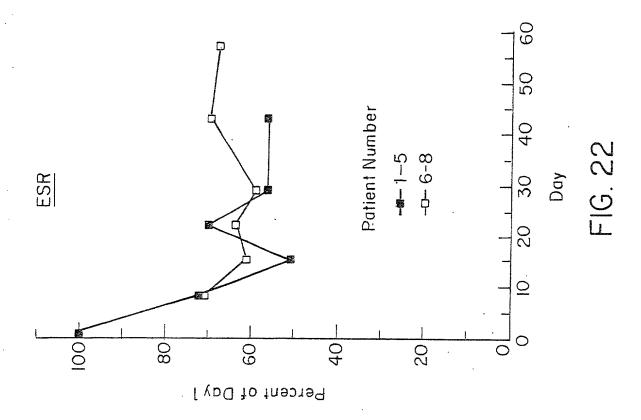


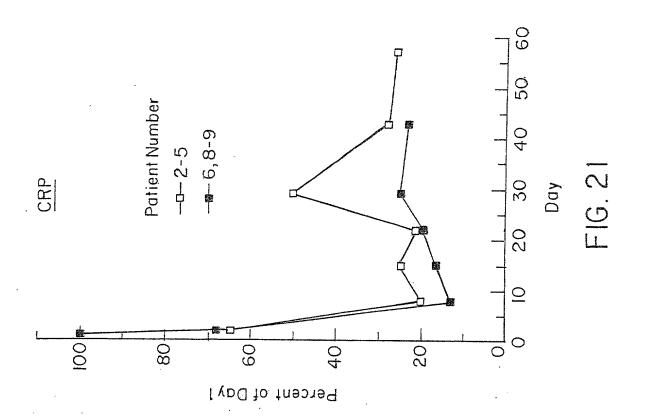
Title: Anti-TNF Antibodies and Peptides...





Title: Anti-TNF Antibodies and Peptides...





Title: Anti-TNF Antibodies and Peptides...

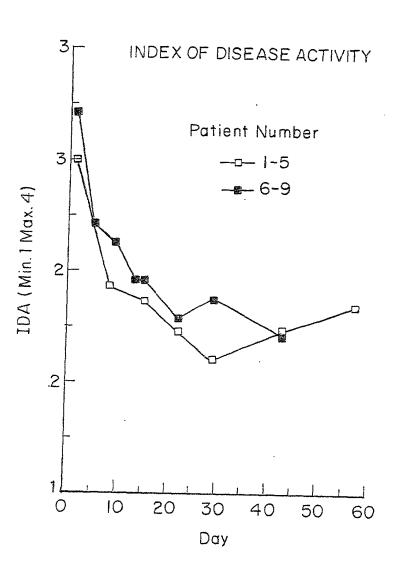
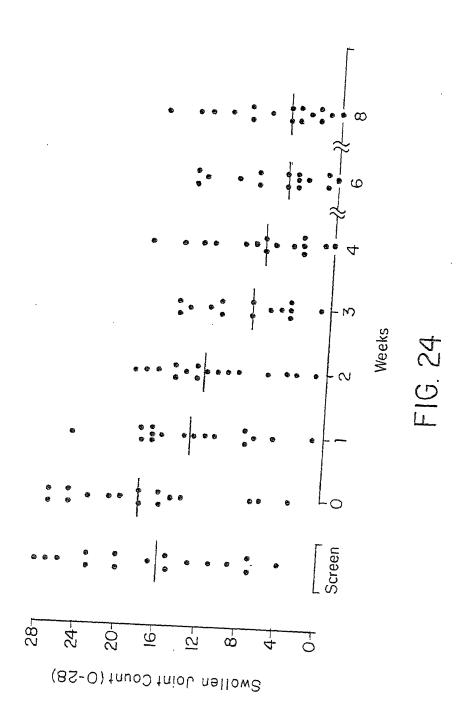
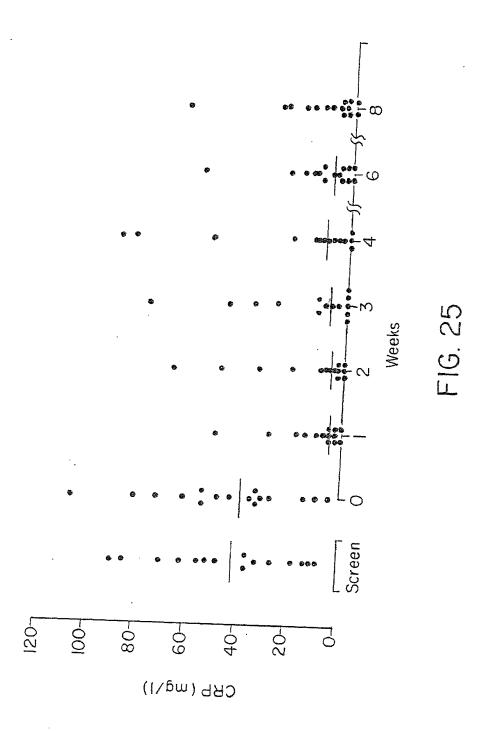


FIG. 23

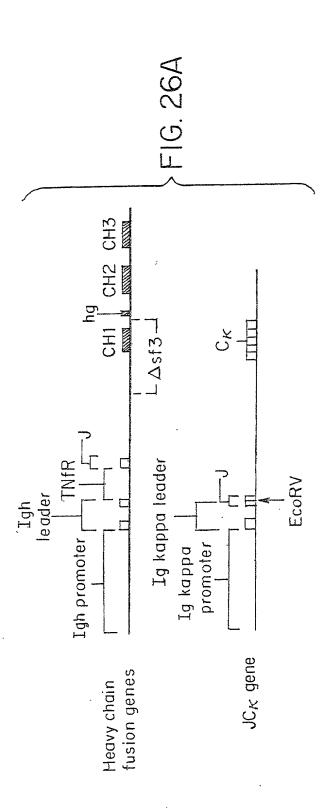
Title: Anti-TNF Antibodies and Peptides...



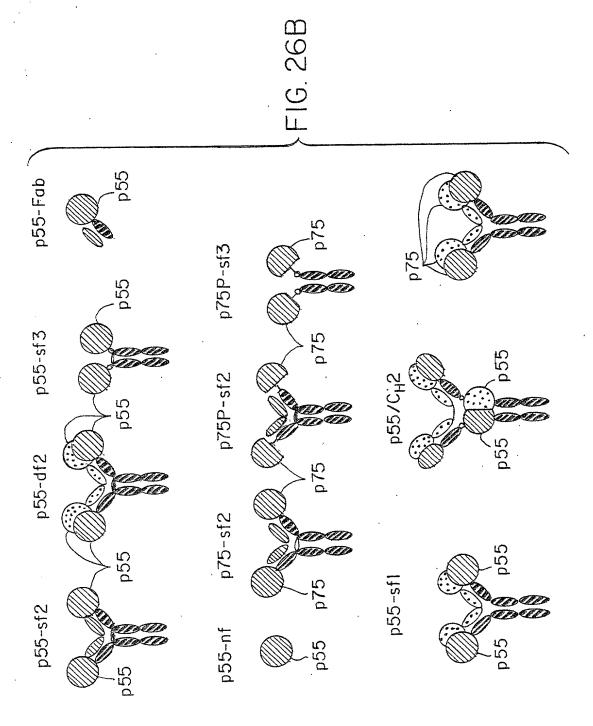
Title: Anti-TNF Antibodies and Peptides...



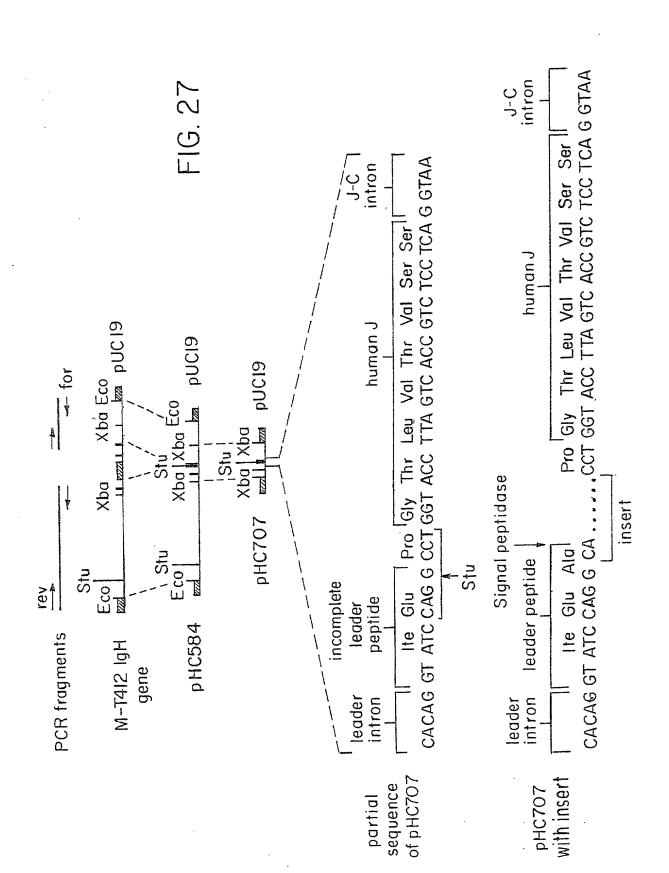
Docket No.: 0975.1005-010
Title: Anti-TNF Antibodies and Peptides...
Inventors: Junming Le, et al.



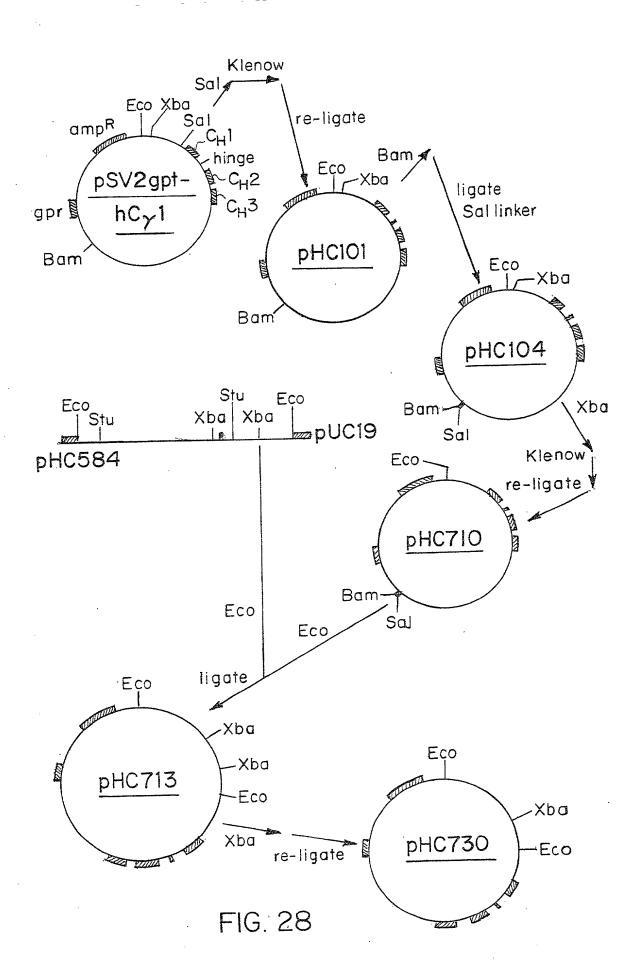
Title: Anti-TNF Antibodies and Peptides...
Inventors: Junming Le, et al.



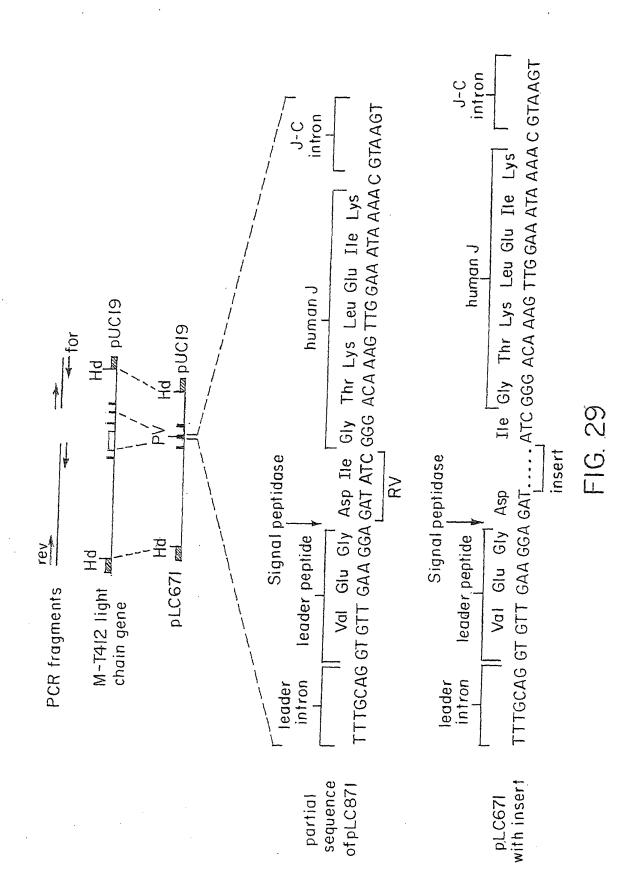
Title: Anti-TNF Antibodies and Peptides...



Title: Anti-TNF Antibodies and Peptides...



Title: Anti-TNF Antibodies and Peptides...



Title: Anti-TNF Antibodies and Peptides...

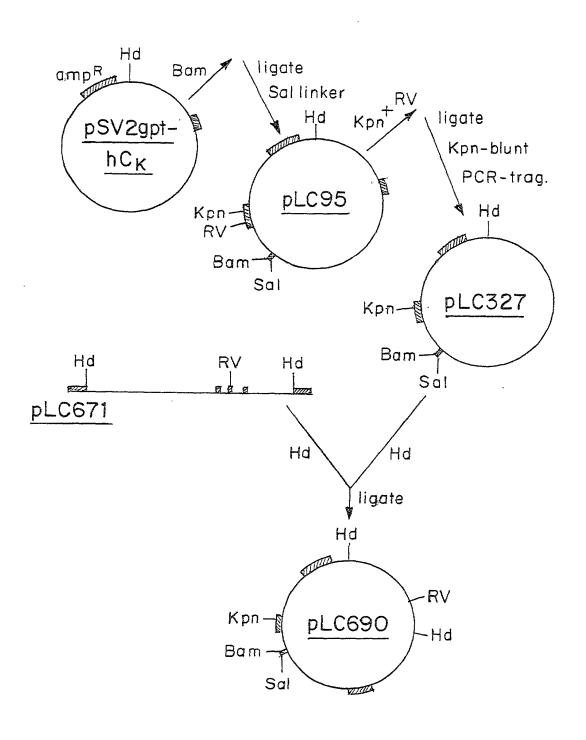


FIG. 30

Title: Anti-TNF Antibodies and Peptides...

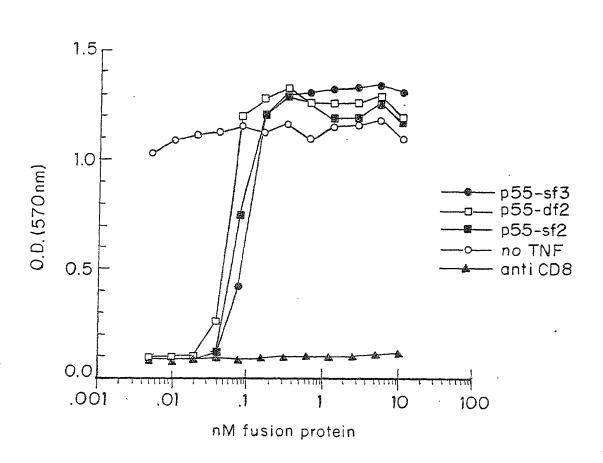


FIG. 31A

Title: Anti-TNF Antibodies and Peptides...

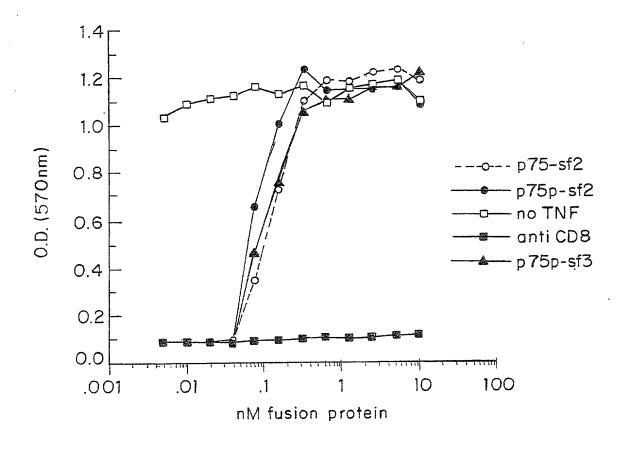


FIG. 31B

Docket No.: 0975.1005-010
Title: Anti-TNF Antibodies and Peptides...
Inventors: Junming Le, et al.

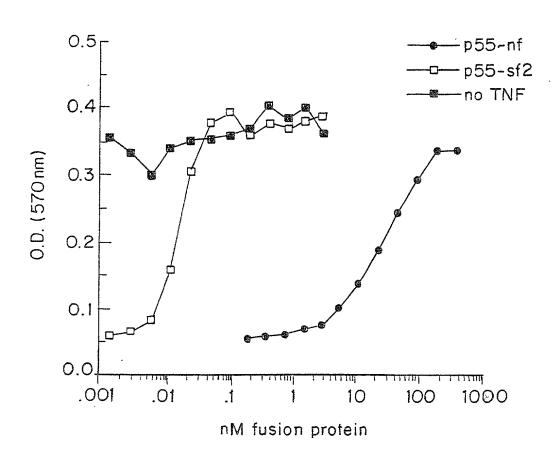


FIG. 31C

Title: Anti-TNF Antibodies and Peptides...

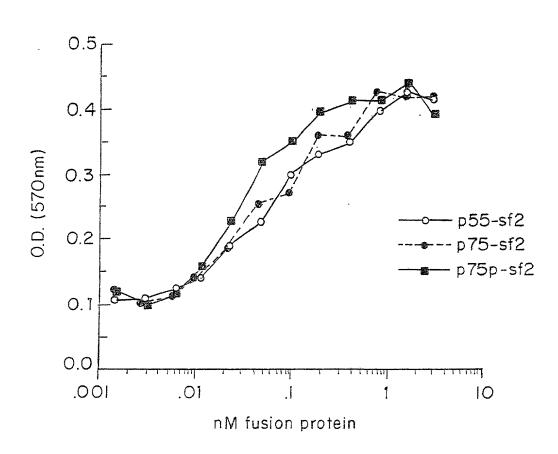


FIG. 32

Title: Anti-TNF Antibodies and Peptides...

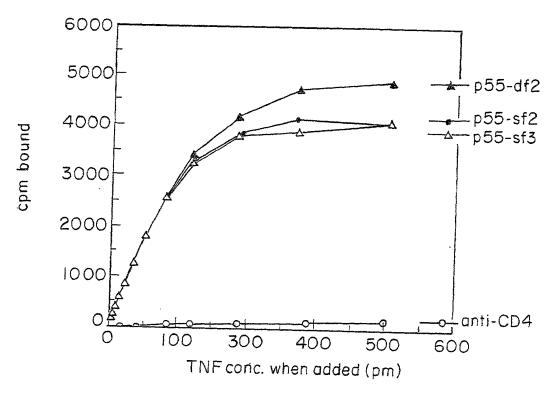


FIG. 33A

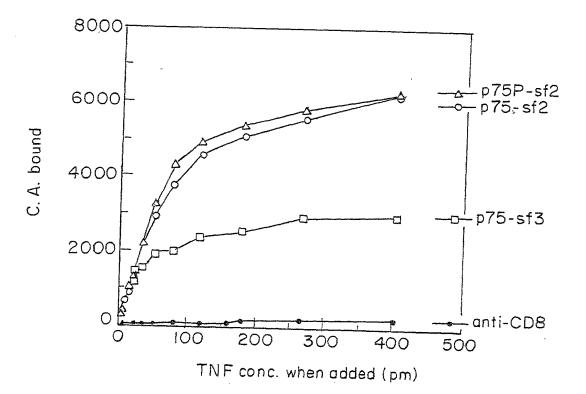
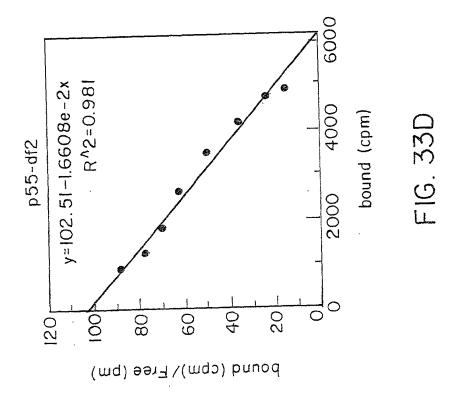
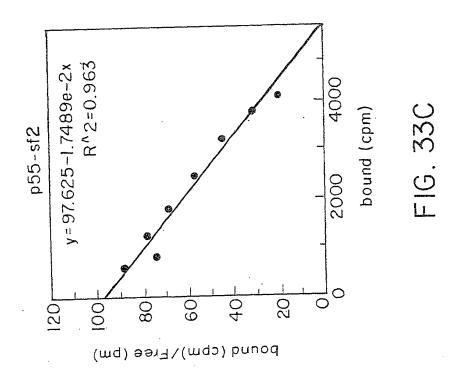


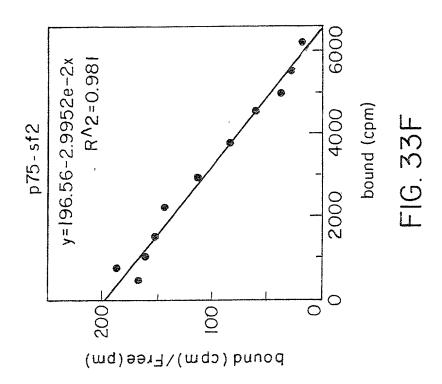
FIG. 33B

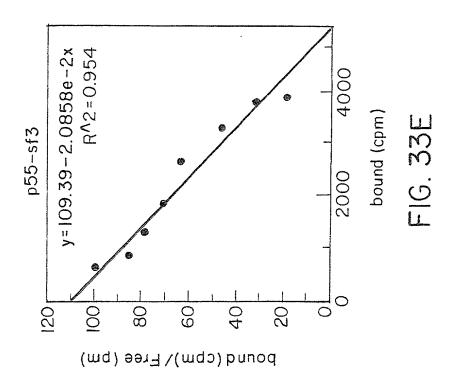
Title: Anti-TNF Antibodies and Peptides...



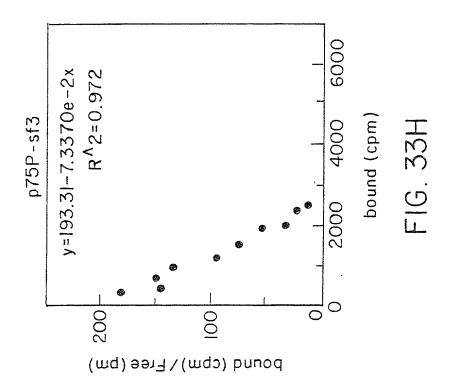


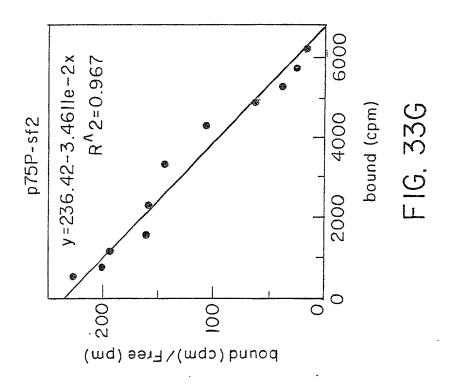
Title: Anti-TNF Antibodies and Peptides...



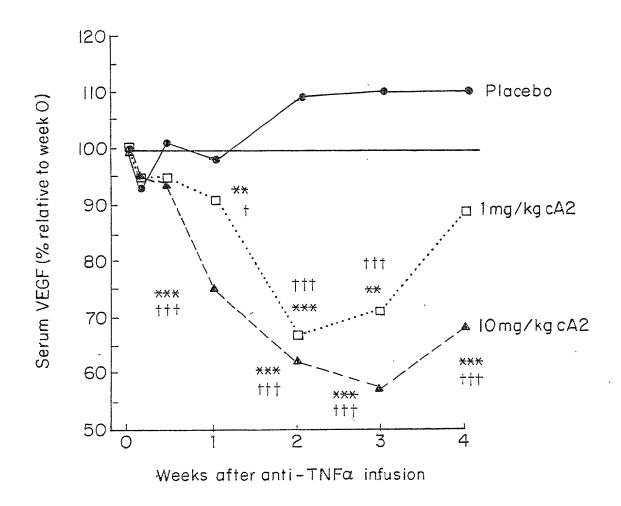


Title: Anti-TNF Antibodies and Peptides...





Title: Anti-TNF Antibodies and Peptides...



* p \leq 0.05, ** p \leq 0.01, *** p \leq 0.001 *versus* pre-infusion † p \leq 0.05, †† p \leq 0.01, †† p \leq 0.001 *versus* change in placebo group

FIG. 34